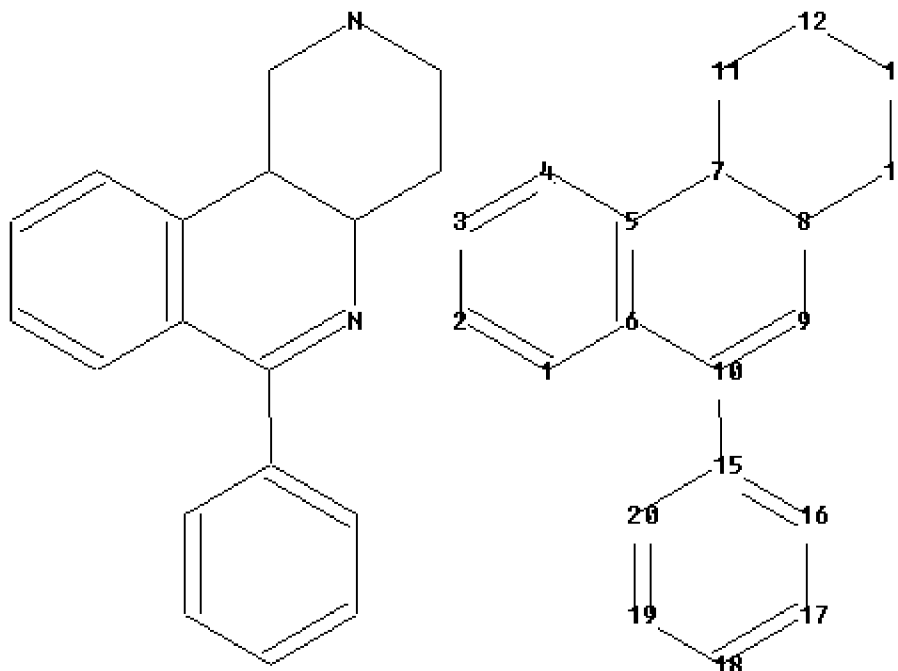


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115 S L2

L4

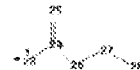
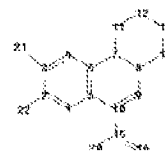
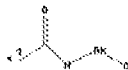
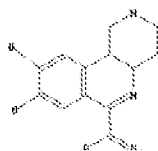
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L6

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L7

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1 S US 20070208051/PN

L8 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2009 ACS on STN

TI Preparation of phenyl benzonaphthyridine derivatives as PDE3/4 inhibitors

ACCESSION NUMBER: 2005:1049863 CAPLUS Full-text

DOCUMENT NUMBER: 143:347067

TITLE: Preparation of phenyl benzonaphthyridine derivatives

as PDE3/4 inhibitors

INVENTOR(S): Kautz, Ulrich; Hatzelmann, Armin; Barsig, Johannes;

Marx, Degenhard; Kley, Hans-Peter; Flockerzi,

Dieter

PATENT ASSIGNEE(S): Altana Pharma A.-G., Germany

SOURCE: PCT Int. Appl., 39 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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EP 1732925	A1	20061220	EP 2005-717070	
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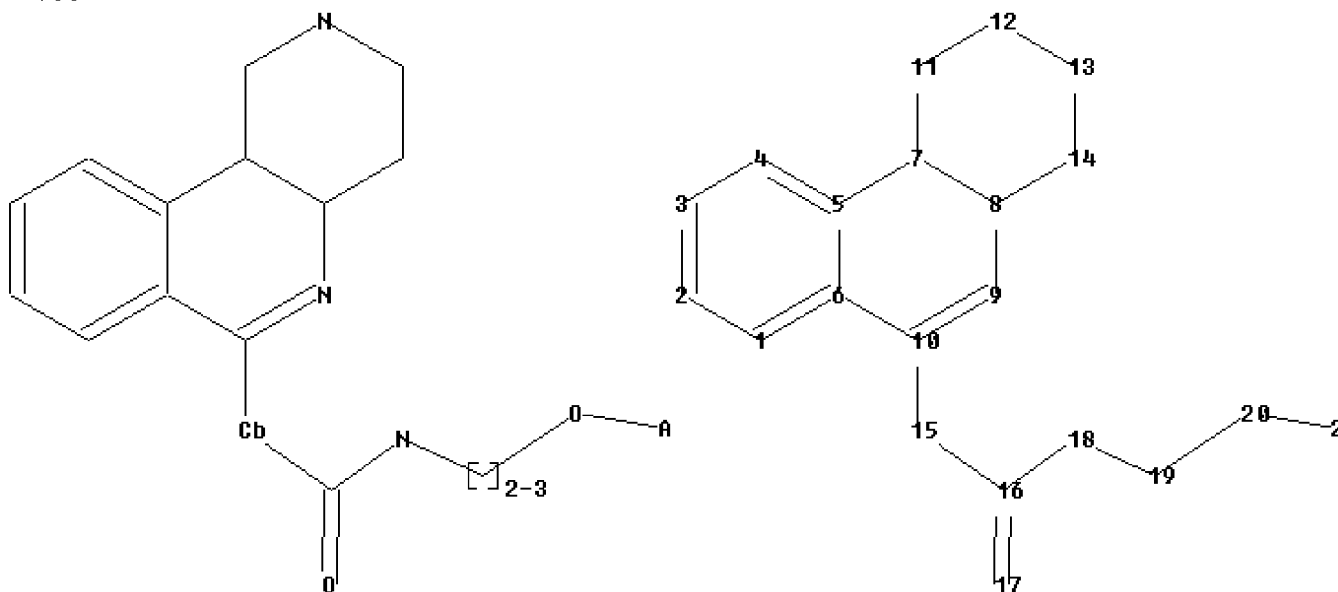
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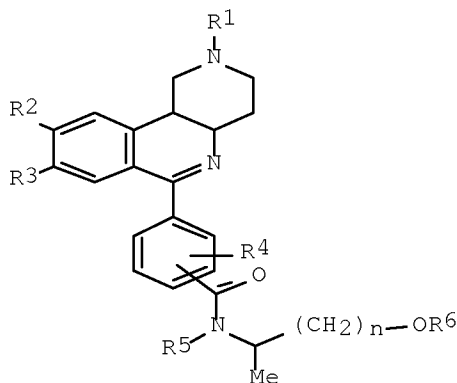
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L9 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2009 ACS on STN  
 TI Preparation of phenylbenzonaphthyridine derivatives as PDE3/4 inhibitors

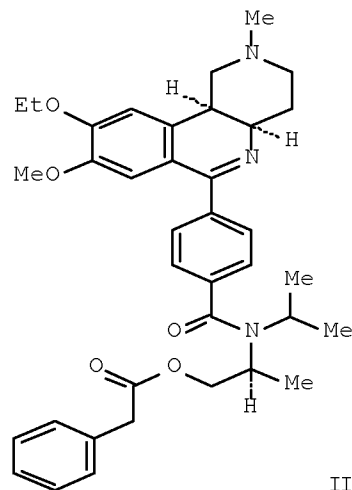
ACCESSION NUMBER: 2004:220332 HCAPLUS Full-text  
 DOCUMENT NUMBER: 140:270839  
 TITLE: Preparation of phenylbenzonaphthyridine derivatives as  
 PDE3/4 inhibitors  
 INVENTOR(S): Flockerzi, Dieter; Hummel, Rolf-peter;  
 Reutter, Felix;  
 Flockerzi, Dieter; Hummel, Rolf-peter;  
 Reutter, Felix  
 PATENT ASSIGNEE(S): Altana Pharma Ag, Germany  
 SOURCE: PCT Int. Appl., 38 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004022557	A1	20040318	WO 2003-EP9617	
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US 20060167034	A1	20060727	US 2005-525566	
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US 7470704	B2	20081230	
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OTHER SOURCE(S):	MARPAT 140:270839		
GI			



I



II

L10 1 S L8 NOT L9

AB Title compds. I [R1 = alkyl; R2 and R3 independently = OH, alkoxy, cycloalkoxy, etc. or R2 and R3 together are alkylenedioxy group; R4 = H, halo, NO2, etc.; R5 = H, alkyl, phenylalkyl, etc.; R6 = alkyl, phenylalkyl or (un)substituted arylalkyl; R7 = alkyl and n = 1-2 or R7 = H and n = 1-3] and their pharmaceutically acceptable salts, are prepared and disclosed as inhibitors of PDE3/4. Thus, e.g., II was prepared by amidation of 4-((4aR,10bS)-9-ethoxy-8-methoxy-2-methyl-1,2,3,4,4a,10b-hexahydrobenzo[c][1,6]naphthyridin-6-yl)benzoic acid (preparation given) with 3-isopropoxypropyl-amine. The inhibitory activity of I towards PDE3 and PDE4 was evaluated using radioactive enzyme assays and it was revealed that compds. of the invention possessed -log IC50 values in the range of 7.8 up to 9.9 mol/L for PDE4 and in the range of 5.8 up to 7.8 mol/L for PDE3. I as inhibitor of PDE3/4 should prove useful in the treatment of respiratory disorders and dermatoses. Pharmaceutical compns. comprising I are disclosed.

ACCESSION NUMBER: 2005:1049863 HCAPLUS Full-text  
DOCUMENT NUMBER: 143:347067

TITLE: Preparation of phenyl benzonaphthyridine  
 derivatives  
 as PDE3/4 inhibitors  
 INVENTOR(S): Kautz, Ulrich; Hatzelmann, Armin; Barsig,  
 Johannes;  
 Marx, Degenhard; Kley, Hans-Peter; Flockerzi,  
 Dieter  
 PATENT ASSIGNEE(S): Altana Pharma A.-G., Germany  
 SOURCE: PCT Int. Appl., 39 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005090345	A1	20050929	WO 2005-EP51204	
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OTHER SOURCE(S): CASREACT 143:347067; MARPAT 143:347067

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L12 0 S L11 NOT L8

FILE 'WPIX' ENTERED AT 14:40:40 ON 09 SEP 2009

L13 0 S L6

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E FLOCKERZI D?/AU

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L15 47 S L14 AND PHOSPHODIESTERASE?

L16 41 S L15 AND (PY<2004 OR AY<2004 OR PRY<2004)

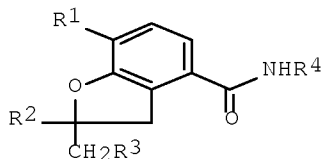
L17 3 S L15 AND (RESPIRATION OR ASTHMA OR BRONCHITIS OR COPD)

L18 1 S L17 AND (PY<2004 OR AY<2004 OR PRY<2004)

L18 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2009 ACS on STN

TI Preparation of phosphodiesterase-inhibiting  
dihydrobenzofurancarboxamides

GI



I

AB The title compds. [I; R1 = (un)substituted alkoxy, cycloalkoxy, PhCH2O, etc.; R2 = alkyl; R3 = H, alkyl; R4 = (un)substituted Ph, pyridyl, etc.], which are potent phosphodiesterase (PDE) inhibitors, useful for the treatment of respiratory diseases [e.g., asthma (no data)] and dermatoses (no data), are prepared Thus, 4-amino-3,5-dichloropyridine was reacted with NaH and 2,3-dihydro-2,2-dimethyl-7-methoxy-4-benzofurancarboxylic acid, producing N-3,5-dichloro-4-pyridyl 2,3-dihydro-2,2-dimethyl-7-methoxy-4-benzofurancarboxamide, m.p. 140-142°, which demonstrated a log IC50 against PDE-IV of 8.47.

IC ICM C07D401-12

ICS C07D307-94; C07D307-79; A61K031-34; A61K031-44

CC 27-16 (Heterocyclic Compounds (One Hetero Atom))

Section cross-reference(s): 1

ST dihydrobenzofurancarboxamide prepn inhibitor phosphodiesterase;

antiasthmatic prepn phosphodiesterase inhibitor  
dihydrobenzofurancarboxamide  
IT Skin, disease  
(dermatoses; phosphodiesterase-inhibiting  
dihydrobenzofurancarboxamides for treatment of)  
IT Respiratory tract  
(disease, phosphodiesterase-inhibiting  
dihydrobenzofurancarboxamides for treatment of)  
IT 177429-18-4P 177429-19-5P 177429-20-8P 177429-21-9P  
177429-22-0P  
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RL: BAC (Biological activity or effector, except adverse); BSU  
(Biological  
study, unclassified); SPN (Synthetic preparation); THU  
(Therapeutic use);  
BIOL (Biological study); PREP (Preparation); USES (Uses)  
(preparation of phosphodiesterase-inhibiting

L18 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2009 ACS on STN

TI Preparation of phosphodiesterase-inhibiting  
dihydrobenzofurancarboxamides

ACCESSION NUMBER: 1996:345409 HCAPLUS Full-text  
DOCUMENT NUMBER: 125:10630  
ORIGINAL REFERENCE NO.: 125:2337a,2340a  
TITLE: Preparation of phosphodiesterase-inhibiting  
dihydrobenzofurancarboxamides  
INVENTOR(S): Amschler, Hermann; Flockerzi, Dieter;  
Gutterer, Beate; Hatzelmann, Armin; Schudt,  
Christian;  
Beume, Rolf; Haefner, Dietrich; Kley, Hans-  
Peter;  
Ulrich, Wolf-Ruediger; Thibaut, Ulrich  
PATENT ASSIGNEE(S): Byk Gulden Lomberg Chemische Fabrik Gmbh,  
Germany  
SOURCE: PCT Int. Appl., 48 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: German  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9603399	A1	19960208	WO 1995-EP2841	
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